Alpha-Iduronidase Enzyme Activity - Mucopolysaccharidosis Type I

Disease Overview

Mucopolysaccharidosis type I (MPS I) is a progressive disorder that ranges in severity and can affect numerous systems throughout the body. Subtypes include attenuated MPS I (previously known as Hurler-Scheie and Scheie syndromes) and severe MPS I (formerly Hurler syndrome). Symptoms vary widely but can include cardiomyopathy, "coarse" facial features (eg, thickening of the earlobes, lips, outside of nostrils, and tongue), intellectual disability, organomegaly, and progressive skeletal dysplasia.1

Genetics

Gene

IDUA

Inheritance

Autosomal recessive

Incidence

MPS I: ~1/70,000

- Attenuated MPS I: 1/500,0001
- Severe MPS I: 1/100,0001

Genotype/Phenotype Correlation

<table>
<thead>
<tr>
<th>MPS I Subtype</th>
<th>Characteristics</th>
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<tbody>
<tr>
<td>Attenuated MPS I</td>
<td>Variants are often milder (eg, single base-pair substitutions)</td>
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<td>- Some residual enzyme function is retained</td>
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<tr>
<td>Severe MPS I</td>
<td>Associated with more severe, nonsense variants</td>
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<td>- Profound alpha-iduronidase deficiency</td>
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Test Interpretation

Results

An extremely low or undetectable level of alpha-iduronidase enzyme activity in leukocytes is consistent with MPS I.

Limitations

Testing for alpha-iduronidase enzyme activity:

- Cannot differentiate between attenuated MPS I (ie, Hurler-Scheie and Scheie syndromes) and severe MPS I (Hurler syndrome)
  - Categorization depends on clinical and/or molecular genetic findings
- Cannot predict carrier status for MPS I
- Does not evaluate enzyme deficiencies in other MPS types

Featured ARUP Testing

Screening

Mucopolysaccharides Screen - Electrophoresis and Quantitation, Urine 0081352
Method: Electrophoresis/Spectrophotometry

Recommended initial test to screen for all mucopolysaccharidosis types, including MPS I

Diagnosis/Monitoring

Alpha-Iduronidase Enzyme Activity in Leukocytes 2011415
Method: Quantitative Fluorometry

Recommended test to exclude MPS I following an abnormal screen or to confirm MPS I in patients with a consistent clinical phenotype and/or a positive family history

Mucopolysaccharidoses Type 1/2, Total Heparan Sulfate and NRE (Sensi-Pro(R)) Quantitative, Urine 3003552
Method: Liquid Chromatography-Tandem Mass Spectrometry

Use to confirm a diagnosis of MPS I or to monitor glycosaminoglycan (GAG) levels in patients with confirmed MPS I

Mucopolysaccharidoses Type 1/2, Total Heparan Sulfate and NRE (Sensi-Pro(R)) Quantitative, Serum or Plasma 3003566
Method: Liquid Chromatography-Tandem Mass Spectrometry

Use to confirm a diagnosis of MPS I or to monitor GAG levels in patients with confirmed MPS I

Mucopolysaccharides, Quantitative, Urine 0081357
Method: Spectrophotometry

Use to monitor GAG levels in patients with confirmed MPS I
Additionally, pseudodeficiency has been demonstrated for this enzyme due to specific gene variants that affect the exogenous substrate but not endogenous substrates. Individuals with pseudodeficiency are not affected with MPS I.

References


Additional Resources


National Organization for Rare Disorders (NORD). Mucopolysaccharidosis type I. [Published: 2019; Accessed: Jan 2022]